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(54) Title: TOPICAL COMPOSITION (57) Abstract <p>The present invention provides topical compositions comprising high surface area zinc oxide, having a surface area between 30 m²/g and 100 m²/g and an average particle size between 0.1 and 200 µm in diameter, said high surface area zinc oxide being present in an amount sufficient to absorb liquids from parts of the body to which the topical composition is applied. The compositions are particularly good at absorbing sweat, sebum, urine and water, making them suitable for treating acne, athletes foot and nappy rash.</p>		

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TOPICAL COMPOSITION

This invention teaches the use of a new form of zinc oxide as an agent to absorb liquids on human skin.

5 The skin is one of the most complex organs in the human body. As part of a myriad of processes, it exudes sweat and sebum. An excess of sweat or sebum is implicated in a number of unpleasant human conditions such as acne, greasy hair/skin and body odour, unless it is removed by washing or some other means.

10 Further a lack of care for the skin, in particular not keeping it dry may result in a number of irritating conditions. Moisture between the toes such as water or sweat provides a fertile breeding ground for the microbes that cause athletes foot. Similarly, the prolonged dampness that results from not changing a baby's nappy frequently or not drying its buttocks sufficiently often gives rise to nappy rash.

15 There are a number of approaches to combating the problems of liquids on the skin.

Where the liquid is sweat, a common approach is to block the sweat pores with antiperspirants. The problem of odour associated with an excess of sweat or sebum being broken down by the micro-organisms present on the skin can
20 be overcome with odour masking agents such as deodorants.

Another approach is to use an agent to absorb the liquid. Talcs, clays and silica have all been used to absorb sebum. Talcs are also used to absorb sweat in foot products, and to keep babies bottoms dry.

25 This invention describes the use of a new form of zinc oxide in topical compositions which has vastly superior absorbence properties. This form of zinc oxide has a surface area between $30\text{m}^2/\text{g}$ and $100\text{m}^2/\text{g}$, preferably greater than $90\text{m}^2/\text{g}$. The particles have an average diameter between 0.1 and $200\mu\text{m}$, preferably between 0.1 and $20.5\mu\text{m}$. Hereinafter, such zinc oxide particles shall be known as "high surface area zinc oxide". The surface area of the material
30 can be determined using laser diffraction. From the surface area, the mean

estimated spherical diameter may be calculated. The diameter of the particles is henceforth taken to mean the mean estimated spheric diameter. The surface area of the high surface area zinc oxide particles used with this invention was calculated using a Micromeritics Flavorsorb II 2300 BET apparatus. Such zinc oxide is commercially available from Elementis as Activox C80.

The production of zinc oxides suitable for use in the topical compositions of the present invention is well documented.

WO 95/04704 (Harcros)

This patent discloses a method for the production of zinc oxide in the form of discrete particles which have an average particle size of 0.08 μm or less in diameter and a surface area $>12.5\text{m}^2/\text{g}$. The zinc oxide produced is found to be particularly good for use as an additive in skin formulations designed for scattering/absorbing UV light.

S. Tichy, SOFW - Journal 119 Jahrgang 8/93

This article teaches a method for producing zinc oxide with a particle size of 200 μm and a surface area of $50\text{-}150\text{m}^2/\text{g}$.

Liu et al - Journal of Materials Science 21 (1986) 3698-3702

This describes another method of producing zinc oxide particles with a diameter of 0.15 μm and a surface area of about $50\text{m}^2/\text{g}$.

These articles are but a small selection from a large number of articles detailing the manufacture of these modified zinc oxide particles.

This invention teaches the use of the high surface area zinc oxide particles to absorb liquids from human skin. Such liquids may include water, urine and in particular sweat and sebum.

Surprisingly, high surface area zinc oxide has been found to be highly efficacious at absorbing liquids, particularly sweat and/or sebum from the skin. Excess liquid upon the skin, especially an excess of sweat and sebum is

implicated as a factor in a number of unpleasant human conditions. These include acne and greasy skin, greasy hair, sweaty feet, athletes foot, body odour and nappy rash.

5 High surface area zinc oxide's high absorbency and gentleness upon the skin make it particularly suitable for use in cosmetic compositions. Cosmetic compositions containing high surface area zinc oxide applied to a part of the body would absorb the liquid present on the skin. Thus, the cosmetic compositions would be capable of acting to treat or prevent many of the socially unacceptable problems which are prevalent in those with an excess of liquid
10 upon their skin.

The present invention provides topical compositions comprising a cosmetically acceptable diluent or carrier and high surface area zinc oxide, having a surface area between $30\text{m}^2/\text{g}$ and $100\text{m}^2/\text{g}$, and an average particle
15 size between 0.1 and $200\mu\text{m}$ in diameter, said high surface area zinc oxide being present in an amount sufficient to absorb liquids from the parts of the body to which the topical composition is applied.

Preferably, the surface area of the high surface area zinc oxide is greater than $90\text{m}^2/\text{g}$ and the average particle size is between 0.1 and $20.5\mu\text{m}$ in
20 diameter.

In preferred topical formulations, the high surface area zinc oxide is present from 1 to 10% , preferably 3 to 8% , most preferably 4 to 6% by weight of the total composition.

25 Acne is a common affliction of many people in their teenage years and sometimes beyond. As a result of puberty, teenagers often have increased levels of sebum. The initial inflammation of the follicle wall in the development of acne results from the presence of free fatty acids derived from the sebum. The normal bacterial flora in the sebaceous duct produce the enzymes responsible for splitting triglycerides in the sebum and releasing these fatty
30 acids. The main micro-organisms in the sebaceous duct are *Propionibacterium acnes* and one or two species of *Staphylococcus aureus*. Therefore in the presence of excess sebum these micro-organisms may result in the development of acne.

Most approaches to a cure for acne focus on trying to absorb the excess sebum or to act on the bacteria present.

5 High surface area zinc oxide particles have been found to be particularly efficacious in the absorption of the straight chain fatty acids found in sebum particularly those of longer chain length. It is these compounds which are particularly implicated in many of the causes of acne. By absorbing these compounds from the skin, high surface area zinc oxide can prevent the formation of pustules and comedones. This invention teaches the use of high surface area zinc oxide in the treatment of acne. High surface area zinc oxide
10 may be used in the preparation of a medicament for the treatment of acne.

Athletes foot is the loose term applied to a skin eruption on the foot, usually between the toes. It is a cutaneous fungal infection, most commonly caused by *Tricophyton rubrum*, *Tricophyton mentagrophytes* or *Epidermophyton floccosum*. In addition to the effect of the micro-organisms, other factors such
15 as wetness or an increase in temperature can contribute to disease development by providing ideal conditions for the initiation and growth of fungal infections. The condition is commonly treated by careful foot hygiene, removing the damp conditions helpful to fungal growth and by the use of antifungal agents. High surface area zinc oxide has been found to be highly absorbent of
20 liquids. By absorbing sweat or liquid, particularly sweat from the skin of the foot, this invention teaches the use of high surface area zinc oxide to treat athletes foot. High surface area zinc oxide may be used in the preparation of a medicament for the treatment of athletes foot.

Nappy rash, the skin eruption which tends to occur on the buttocks of
25 infants is due to infrequent changing of soiled nappies. The condition is often worsened by secondary infection with *Candida albicans*. To prevent this condition, nappies are changed regularly and care is taken to ensure that the baby's bottom is dried properly. The buttocks are commonly treated with an agent to absorb any surplus liquid. High surface area zinc oxide has been
30 found to be highly efficacious in absorbing liquid on the skin in this region. This invention teaches the use of high surface area zinc oxide to prevent and treat nappy rash. High surface area zinc oxide may be used in the preparation of a medicament for the treatment of nappy rash.

In an embodiment of the invention, the cosmetic formulation may be suitable for application as an deodorant. The deodorant may be in the form of a roll on, a spray or other suitable form. Such compositions may be formulated in a manner known to those skilled in the art, and may include, though not limited to:-

5

a) antiperspirants such as aluminium zirconium, aluminium chlorohydrate, aluminium zirconium pentachlorohydrate.

b) emulsifiers such as steareth-2, steareth-21 and polypropylene glycol-15 stearyl ether.

10

c) thickeners such as cyclomethicone, dimethicone copolyol, hydroxypropyl methylcellulose, hydroxypropylcellulose.

d) humectants such as propylene glycol, butylene glycol and glycerin.

In a further embodiment of the invention, the cosmetic formulation may be a shampoo. The composition may be formulated in a manner known to those skilled in the art. Such compositions may include, though not limited to:-

15

a) surfactants such as cocamidopropyl betaine and sodium laureth sulphate.

b) thickeners such as xanthan gum or hydroxyethylcellulose, laureth-3, polyethylene glycol-40 hydrogenated castor oil, polyethylene glycol-55 propylene glycol oleate and propylene glycol.

20

c) pearl concentrates such as formaldehyde, a blend of stearic acid, cocamide MEA, glycol distearate and glycol stearate, methyldibromo glutaronitrile.

d) conditioners such as polyquaternium-39 and polyquaternium-7, hydroxypropyl guar hydroxypropyltrimonium chloride, polyquaternium-10.

25

e) preservatives such as paraben, phenoxyethanol with methyldibromo glutaronitrile.

f) perfumes.

g) colour.

5 In another embodiment of the invention, the cosmetic formulation may be a gel. The composition may be formulated in a manner known to those skilled in the art. Such compositions may include, though not limited to:-

a) antibacterials such as dichlorobenzyl alcohol, triclosan, chlorhexidine digluconate and salicylic acid.

10

b) alcohols such as denatured ethanol and isopropyl alcohol.

c) humectants such as panthenol, butylene glycol, glycerin and propylene glycol.

15

d) preservatives such as methyldibromo glutaronitrite, phenoxyethanol, magnesium chloride, magnesium nitrate, methylchloroisothiazoline, any paraben and methylisothiazolinone.

20

e) solubilisers such as polysorbate 20, polypropylene glycol-40 hydrogenated castor oil.

f) cooling agents such as Hamamelis virginiana solution and Mentha piperita.

25

h) emollients such as glycerin, propylene glycol and butylene glycol.

i) oil absorbers such as silica.

30

j) thickeners such as xanthan gum, hydroxyethylcellulose, sodium magnesium silicate.

In another embodiment of the invention, the cosmetic formulation may be a powder, such as, but not limited to, foot powder or face powder. The

composition may be formulated in a manner known to those skilled in the art. Such compositions may include, though not limited to:-

- a) preservative such as any paraben, such as methyl paraben, ethyl paraben, propyl paraben and butyl paraben.
- 5 b) anti-caking agent/lubricants such as magnesium stearate, calcium stearate and stearic acid.
- c) binders such as paraffinum liquidum and any waxes and oils.
- d) powders such as sanitised talc.
- 10 f) colours
- g) absorbents such as Zea mays, rice starch and sodium bicarbonate.
- 15 h) flow agents such as silica.

In a further embodiment of the invention, the cosmetic formulation may be a skin wash, such as a cleanser, moisturiser, face wash, lotion, stick or cream. The composition may be formulated in a manner known to those skilled in the art. Such compositions may include, though not limited to:-

- a) alcohols such as isopropyl myristate, stearyl alcohol, denatured ethanol.
- b) emulsifiers such as steareth-2, glyceryl stearate, hydrogenated vegetable glycerides, steareth-21, ceteth-20, cetyl alcohol, cetearyl alcohol, stearic acid, paraffin, stearyl alcohol, polawax, tribehenin, cetareth-7, ceteth-5.
- 25 c) emollients such as polypropylene glycol-5-ceteth-20, methyl gluceth-10, dicaprylyl maleate, cetearyl isononanoate, silicones, paraffinum liquidum, octyl palmitate, petrolatum, dioctyl maleate, isohexadecane, cetearyl octanoate and isopropyl myristate.
- d) solubilisers such as polysorbate 80, polysorbate 20, polyethylene glycol-40
30 hydrogenated castor oil, any polysorbate.

- e) antibacterials such as triclosan, chlorhexidine digluconate, salicylic acid, dichlorobenzyl alcohol.
- g) thickeners such as hydroxyethylcellulose, xanthan gum, sodium magnesium silicate, magnesium aluminium silicate, cellulose.
- 5 i) detergents such as sodium laureth sulfate, ammonium lauryl sulfate, magnesium lauryl sulfate, disodium undecylenamido MEA-sulfosuccinate.
- j) preservatives such as phenoxyethanol, 2-bromo-2-nitropropane-1,3-diol, methyl dibromo glutaronitrile, imidazolidinyl urea, magnesium chloride,
10 magnesium nitrate, methylchloroisothiazolinone, methylisothiazolinone or any paraben, such as butyl paraben, ethyl paraben, methyl paraben and propyl paraben.
- j) absorbents such as hydrated silica, clays, talcs.
- 15 k) antioxidants such as butylated hydroxytoluene or butylated hydroxyacetone.
- l) moisturisers such as butylene glycol, propylene glycol, sorbitol and glycerin, panthenol, sodium hyaluronate, sodium PCA.

20

The efficacy of high surface area zinc oxide as an absorbent has been demonstrated by *in vivo* and *in vitro* trials. The suitability of high surface area zinc oxide for the purposes stated herein has been demonstrated by trials of standard control formulations against those formulations containing high surface
25 area zinc oxide.

25

The invention is further illustrated by way of the following non-limiting examples:-

30

Unless otherwise stated, the zinc oxide used in the following examples has an average surface area of 90m²/g and an average particle diameter of 10.47 μ m.

Example 1 - Foot Powder + High Surface Area Zinc Oxide

	<u>Ingredient</u>	<u>%</u>
	Sanitised talc	47.46
	High Surface Area Zinc Oxide	5.00
5	Zea Mays	47.46
	Silica	0.08

10 Silica and half the sanitised talc were sifted through a 40 mesh sieve into a bowl containing the high surface area zinc oxide, then mixed until thoroughly dispersed.

15 The remaining sanitised talc was sifted into a Simon Solitec Mixer via a 20 mesh sieve. To this was added the high surface area zinc oxide mixture, via a 40 mesh sieve. The Zea Mays was added and the mixture stirred for 10 minutes. A fifth of the mixture was then removed and sifted back into the mixer via a 20 mesh sieve. The mixture was then stirred for a further 20 minutes.

Example 2 – Foot Powder Control

20	<u>Ingredient</u>	<u>%</u>
	Sanitised talc	50.00
	Zea Mays	50.00

25 Ingredients were sieved through a 40 mesh sieve and then mixed for 20 minutes.

Example 3 – Foot Powder with High Surface Area Zinc Oxide

30	<u>Ingredient</u>	<u>%</u>
	Sanitised Talc	47.5
	Zea Mays	47.5
	High surface area zinc oxide	5.0

35 Ingredients were sieved through a 40 mesh sieve and then mixed for 20 minutes.

Example 4 - Roll-on Deodorant + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
	High Surface Area Zinc Oxide	5.00
5	Polypropylene glycol-15 stearyl ether	4.00
	Steareth-21	1.50
	Steareth-2	2.20
	Aluminium Chlorohydrate Solution	30.00
	Tetrasodium EDTA (Sequestrene)	0.10
10	Perfume	0.50
	Colour	qs
	Purified Water	to 100

15 Tetrasodium EDTA was added to purified water, mixed with a homogeniser for 1 minute, then warmed to 70-75°C, until all the solids had dissolved.

20 Polypropylene glycol - 15 stearyl ether, steareth-21, steareth-2 and high surface area zinc oxide were mixed and heated to 70-75°C prior to addition of tetrasodium EDTA in water. The mixture was homogenised for 5 minutes then stirred slowly whilst cooling to 35°C. Then the aluminium chlorohydrate solution was slowly added to the stirred solution, followed by the perfume. The mixture was stirred until homogeneous, then the colour solution was added.

Example 5 - Oily Scalp Shampoo + High Surface Area Zinc Oxide

25	<u>Ingredient</u>	<u>%</u>
	Panthenol	0.20
	Xanthan Gum	0.40
	Sodium Laureth Sulfate	35.00
30	Cocamidopropyl betaine 30 %	1.50
	Citric acid	0.08
	Polyquaternium - 39	0.50
	Laureth - 3	1.00
	Silk protein complex	0.05
35	Preservative	0.20
	Perfume	0.50
	Colour	qs

	Salt	1.62
	Pearling agent blend consisting of:	
	Formaldehyde, glycol distearate, laureth - 10,	
	cocamide MEA and sodium laureth sulfate	4.00
5	Purified Water	52.95
	High Surface Area Zinc Oxide	2

High surface area zinc oxide was added to a solution of citric acid in water, and mixed for 15 minutes. Xanthan gum was then added and the mixture homogenised for 15 minutes. To the mixture was added sodium laureth sulfate, cocamidopropyl betaine 30%, laureth-3, panthenol, polyquaternium-39, silk protein complex, perfume and preservative and the pearling agent blend whilst stirring. Colour solution was added and then cold water to make up to bulk. The mixture was then stirred until uniform.

Example 6 - Mens Facial Wash + High Surface Area Zinc Oxide

	<u>Ingredient</u>	<u>%</u>
	High Surface Area Zinc Oxide	5.00
20	Hydroxyethylcellulose	1.25
	Sodium laureth sulfate	6.57
	Disodium undecylenamido sulfosuccinate	1.00
	Butylene Glycol	2.00
	Preservative	0.80
25	Benzoic Acid	0.10
	Polysorbate 20	2.00
	Perfume	0.40
	Herbal extract	0.60
	Colours	qs
30	Purified Water	to 100

Stage 1

In a stainless steel vessel butylene glycol and preservative were mixed together until uniform.

Stage 2

In another container the perfume and polysorbate 20 and high surface area zinc oxide were mixed together until uniform.

Stage 3

5 To a stainless steel container was added some of the water and hydroxyethylcellulose. The mixture was stirred for 20-30 minutes until fully dispersed.

10 Stage 3 was then added to stage 1, followed by the herbal extract, benzoic acid and colour. The mixture was stirred until fully dispersed, then sodium laureth sulfate, and sodium undecylenamido MEA-sulfosuccinate were added to the stirred mixture. Stage 2 was added and mixed thoroughly. Cold water was added to make up to bulk.

15 Example 7 - Skin Treatment Gel + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
	Alcohol denatured	10.00
	Allantoin	0.10
20	Glycerin	1.00
	Butylene Glycol	4.00
	Xanthan Gum	1.00
	Phenoxyethanol	0.20
	Hydrated silica	0.50
25	Dichlorobenzyl alcohol	0.10
	Colour	qs
	Benzophenone - 4	0.10
	Purified Water	to 100
	Panthenol	0.50
30	High Surface Area Zinc Oxide	5.00

Xanthan gum, dispersed in 2% of the butylene glycol was added to some of the purified water, and mixed together for 30 minutes. To this mixture allantoin, sequestrene and panthenol were added and the mixture stirred for 5 minutes.

35 The mixture was cooled to 35°C, then premixed phenoxyethanol and glycerin were added to the mixture, followed by premixed alcohol (denatured) and

purified water followed by premixed dichlorobenzyl alcohol and butylene glycol. The mixture was then stirred. Benzophenone-4 and water was then added with stirring, followed by hydrated silica and high surface area zinc oxide. The mixture was stirred, cooled to below 35°C and then the colour was added. Cold water was added to make up to bulk, and the mixture stirred for a further 30 minutes.

Example 8 - Non-oily Moisturiser + High Surface Area Zinc Oxide

10	<u>Ingredients</u>	<u>%</u>
	High Surface Area Zinc Oxide	5.00
	Colour	qs
	Perfume	0.10
	Triclosan	0.10
15	Allantoin	0.10
	Phenoxyethanol	0.20
	Hydroxyethylcellulose	2.00
	Polysorbate 20	1.00
	Butylene Glycol	3.50
20	Glycerin	4.50
	Purified Water	to 100

Stage 1

To purified water in a homogeniser hydroxyethylcellulose was added and then homogenised for at least 30 minutes. The homogeniser was switched off and with stirring allantoin and phenoxyethanol, which had been previously dissolved in glycerin and butylene glycol, were added.

Stage 2

Butylene glycol and glycerin were warmed together to 45°C. Then with stirring triclosan and high surface area zinc oxide were added and completely dissolved and cooled to 35°C.

Stage 3

Using a homogeniser stage 2 was added to stage 1 and homogenised for 10 minutes. The perfume, previously dispersed in polysorbate 20 was then added and stirred in well. The colour was added and the emulsion was then

homogenised for a further 5 to 10 minutes until the product was smooth. Purified water, sufficient to make it up to bulk was then added.

Example 9 - Cleansing Lotion + High Surface Area Zinc Oxide

5

<u>Ingredients</u>		<u>%</u>
High Surface Area Zinc Oxide		5.00
Purified Water		to 100
Colour		qs
10	Melaleuca alternifolia	0.50
	Polypropylene glycol-5-ceteth-20	3.25
	Polysorbate 80	0.20
	Citric acid	0.12
	Disodium Phosphate	0.38
15	Triclosan	0.30
	Butylene Glycol	0.20
	Alcohol (denatured)	48.00

20

Stage 1

Alcohol (denatured) and triclosan were mixed together until homogeneous. Water was added and mixed well. Butylene glycol was then added and the mixture stirred.

25

Stage 2

To a suitable stainless steel container disodium phosphate and water were added, and warmed to 55-60°C with stirring. Then more water was added, with stirring and the solution was then allowed to cool.

30

Stage 3

35

When the temperature of stage 2 had reached 20-25°C it was added to stage 1 with stirring. Citric acid was added and mixed well. In a suitable stainless steel container some of the polypropylene glycol-5-ceteth-20, the melaleuca alternifolia and high surface area zinc oxide were added and mixed thoroughly and then added to the main vessel. In a suitable container the remaining polypropylene glycol-5-ceteth-20 and polysorbate 80 which was then

added to the main vessel. Colour was added, followed by water to make up to bulk.

Example 10 – Toner and Cleanser + High Surface Area Zinc Oxide

5	<u>Ingredients</u>	<u>%</u>
	Purified water	43.30
	Citric acid	0.12
	Sodium citrate	0.38
10	Herbal extract	1.00
	Glycerin	2.00
	Polysorbate 80	0.20
	Alcohol denatured	48.00
	High surface area zinc oxide	5.00

15

The ingredients were added together and stirred until fully dissolved.

Example 11 – Light Moisture Fluid + High Surface Area Zinc Oxide

20	<u>Ingredients</u>	<u>%</u>
	Pentaerythrityl tetraisostearate	4.00
	Xanthan gum	0.25
	Tetrasodium EDTA	0.10
	Glycerin	1.00
25	Butylene glycol	2.00
	Methylparaben	0.20
	Polyethylene ester-2-stearyl alcohol	2.00
	Polyethylene ester-21-stearyl alcohol	1.00
	Cetyl alcohol	2.50
30	Hydroxyethyl cellulose	0.40
	Cetearyl isononanoate	2.00
	Purified water	76.84
	Butylated hydroxytoluene	0.02
	Butyl methoxydibenzoylmethane	0.50
35	Octyl methoxycinnamate	0.99

	2-Bromo-2-nitropropane-1,3-diol	0.03
	Citric acid	0.06
	Sodium citrate	0.11
	Silk powder	0.50
5	Herbal extract	0.40
	Propylparaben	0.10
	High surface area zinc oxide	5.00

10 Hydroxyethylcellulose was dispersed into an aqueous solution of tetrasodium EDTA. Xanthan gum in glycerin was added to the mixture, followed by methyl paraben. The mixture was stirred and warmed to 70-75°C to create the water phase.

15 Cetyl alcohol, high surface area zinc oxide, polyethylene ester-2-stearyl alcohol, polyethylene ester-21-stearyl alcohol, propyl paraben, cetearyl isononanoate, pentaerythrityl tetraisostearate, butylated hydroxy toluene, butyl methoxydibenzoyl methane and octyl methoxycinnamate were mixed together and warmed to 70-75°C to create the oil phase.

20 The oil and water phases were then emulsified together for 5-10 minutes before being force cooled to 30-35°C. An aqueous solution of citric acid, sodium citrate and silk powder was added to the emulsion. Herbal extracts were then added, and the emulsion made up to bulk with water.

25 **Example 12 - Cleansing Wash + High Surface Area Zinc Oxide**

	<u>Ingredients</u>	<u>%</u>
	Cocamidopropyl betaine 30%	5.00
	Benzophenone - 4	0.10
30	Sodium citrate	0.60
	Disodium undecylenamido MEA-sulfosuccinate solution	1.00
	Triclosan	0.20
	Sodium chloride	1.00

	Laureth - 3	2.00
	Sodium laureth sulfate	47.20
	Melaleuca alternifolia	0.500
	Phenoxyethanol	0.15
5	Colour	qs
	Citric acid	0.10
	Purified Water	to 100
	High Surface Area Zinc Oxide	5.00
10	Stage 1	
	In a vessel sodium laureth sulfate, melaleuca alternifolia and high surface area zinc oxide were mixed until uniform.	
	Stage 2	
15	In the base pan the triclosan was dispersed in cocamidopropyl betaine 30% and stirred for 5 minutes. The purified water was added and stirred well. Citric acid was added and stirred until dissolved followed by sodium citrate, which was stirred until dissolved. The mixture was then stirred for a further 10 minutes and subsequently cooled to 30-35°C. Premixed phenoxyethanol in water, followed by	
20	benzophenone-4 in water were then added. Colour was added followed by saline solution, then water was added to make up to bulk.	

Example 13 - Moisture Fluid + High Surface Area Zinc Oxide

25	<u>Ingredients</u>	<u>%</u>
	High Surface Area Zinc Oxide	5.00
	Glyceryl stearate	1.50
	Steareth-2	2.00
	Steareth-21	1.00
30	Cetyl Alcohol	1.00
	Glycerin	1.00
	Butylene Glycol	2.00
	Purified Water	86.50

35 Stage 1

Glyceryl stearate, steareth-2, steareth-21, cetyl alcohol and high surface area zinc oxide were melted together at 70-75°C.

Stage 2

Glycerin was dissolved with stirring in water at 70-75°C.

5 Stage 3

Stage 1 was then added to stage 2 with stirring, then homogenised for 15 minutes. Water was added to the stirred mixture, then the mixture was cooled to 35°C. Butylene glycol was added and the mixture stirred until homogeneous, then made up to bulk with water.

10

Example 14 - Pressed Powder + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
	High Surface Area Zinc Oxide	5.00
15	Sanitised talc	85.72
	Magnesium Stearate	5.00
	Methylparaben	0.10
	Red colour	0.23
	Yellow colour	0.45
20	Paraffinum liquidum	2.10
	Petrolatum	1.40

Sanitised talc, magnesium stearate, methylparaben, high surface area zinc oxide and colours were mixed together for 10 minutes at high speed.

25 Paraffinum liquidum and petrolatum were mixed together, heated to 75°C and then sprayed into the bulk mixture at low speed. The bulk was mixed for 5 minutes, then passed twice through a hammer mill before being passed through a 30 mesh seive.

30 **Example 15 - Cover Up Stick + High Surface Area Zinc Oxide**

	<u>Ingredient</u>	<u>%</u>
	Chalk	23.31
	Carnauba	1.29
35	Candelilla Cera	1.01
	Hydrocarbon consisting of Cera Microcristallina, paraffin and polyethylene	4.60

	Cera Microcristallina	4.22
	Butylated hydroxyacetone	0.03
	Propylparaben	0.10
	Octyldodecanol	46.02
5	Triclosan	0.19
	Allantoin	0.14
	Pigment	11.98
	High Surface Area Zinc Oxide	5.00
	Synthetic wax	2.01

10

Pigments, high surface area zinc oxide and chalk were added to the Diosna mixer and mixed for 30 minutes. The mix was then passed through the Mikro mill, then a vibrating sieve to give the colour preparation.

15

White wax, Carnauba and candelilla cera were added to a stainless steel steam jacketed pan fitted with a premier dispersator head, and melted together at 90-95°C. To the melt was added the hydrocarbon wax. When melted, octyldodecanol was added and the mixture stirred.

20

The mixture was cooled to 85-90°C then propylparaben, butylated hydroxyacetone and triclosan were added to the stirred mixture, followed by allantoin and then by the colour preparation. The mixture was then stirred for a further ten minutes.

The mixture was then stirred through a 40 mesh sieve into a shallow tray and stirred slowly until set.

25

Example 16 - Shower Gel + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
	Purified Water	47.48
	Perfume	0.10
30	Colour	0.00075
	Sodium chloride	1.19
	Dichlorobenzyl Alcohol	0.50
	Butylated Hydroxytoluene	0.0048

	Triclosan	0.2916
	PEG – 7 Glyceryl Cocoate	2.916
	Citric Acid	0.0216
5	A preservative blend consisting of: Phenoxyethanol, Butylparaben, Ethylparaben, Methylparaben and Propylparaben	0.80
	Cocamidopropyl Betaine	5.83
	Sodium Laureth Sulfate	45.89
10	High Surface Area Zinc Oxide	5.00

High surface area zinc oxide was added to purified water and mixed well. Sodium chloride and citric acid were then added and the mixture stirred until both had dissolved. Sodium laureth sulfate, cocamidopropyl betaine, phenoxyethanol,
15 butylparaben, ethylparaben, methylparaben and propylparaben were then added and the mixture stirred.

PEG 7 Glyceryl cocoate, butylated hydroxytoluene, triclosan and dichlorobenzyl alcohol were mixed together and warmed to 45°C. Perfume was then added and
20 the mixture stirred until homogenous.

The two mixtures were then combined and stirred until homogenous. Colour solution was added to the stirred mixture followed by water to make up to bulk. The mixture was then stirred until uniform.

25

Example 17 – Acne Lotion + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
	Purified Water	85.115
30	Benzoyl Peroxide	6.667
	Hydroxyethylcellulose	1.00
	Citric Acid	1.53
	Sodium Hydroxide	0.6915

High Surface Area Zinc Oxide

5.00

Sodium hydroxide was added to a stirred aqueous solution of citric acid. Hydroxyethylcellulose was then added to the mixture which was then stirred for 30 minutes. Benzoyl peroxide was then added followed by some water. The mix was stirred for 2 minutes then homogenised for 20 minutes under vacuum. High surface area zinc oxide was then added to the mixture and stirred thoroughly. Aqueous sodium hydroxide was then added to the stirred mixture, which was then stirred for a further hour.

10

Example 18 – Sensitive Cleansing Pads + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
	Purified Water	68.53
15	A preservative blend consisting of: Phenoxyethanol Butylparaben, Ethylparaben, Methylparaben and Propylparaben	0.80
	Perfume	0.05
	Ethoxylated (20) sorbitan monolaurate	1.02
20	Sodium Citrate	0.21
	Citric Acid	0.06
	Cetrimoniumbromide	0.51
	Alcohol (denatured)	17.38
	PPG – 5 – Ceteth – 20	0.31
25	Butylene Glycol	4.60
	Glycerin	1.02
	Chlorhexidine digluconate	0.51
	High Surface Area Zinc Oxide	5.00

30 Chlorhexidine digluconate, butylene glycol, PPG-5-Ceteth-20, glycerin, cetrimoniumbromide, citric acid, sodium citrate were dissolved in purified water. Denatured alcohol, high surface area zinc oxide and the preservative blend were then added and the mixture stirred. A mixture of ethoxylated (2) sorbitan

monolaurate and perfume was then added and the mixture stirred. Purified water was added to make up to bulk. The mixture was stirred for 30 minutes then pumped through an 80 mesh sieve to a suitable storage vessel. Rayon/polyester pads were then impregnated with the mixture to give the sensitive cleaning pads.

5

Example 19 – Overnight Gel + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
	Purified Water	50.6
10	Perfume	0.05
	Triclosan	0.10
	Dichlorobenzyl Alcohol	0.50
	Alcohol (denatured)	39.50
	Hydroxyethylcellulose	1.25
15	Glycerin	3.00
	High Surface Area Zinc Oxide	5.00

Hydroxyethylcellulose and glycerin were dispersed in water then transferred to a mixer via a sieve covered with muslim.

20

Denatured alcohol, dichlorobenzyl alcohol and triclosan were mixed together until homogenous. Perfume and then high surface area zinc oxide were then dispersed in the mixture. The mixture was then transferred to the mixer under vacuum via a sieve. The mixture was stirred for 30 minutes until homogeneous. Water was then

25

Example 20 – Emergency Gel + High Surface Area Zinc Oxide

	<u>Ingredients</u>	<u>%</u>
30	Purified Water	37.00
	Hydroxypropyl Methylcellulose	2.50
	Sodium Citrate	0.30
	Alcohol (denatured)	20.00

	Butylene Glycol	15.00
	Propylene Glycol	18.00
	Triclosan	0.20
	Salicylic Acid	2.00
5	High Surface Area Zinc Oxide	5.00

Propylene glycol, butylene glycol and ethanol were mixed together. Salicylic acid and triclosan were then dissolved in the mixture. High surface area zinc oxide was then added to the stirred mixture. Hydroxypropyl methylcellulose was then dispersed in the mixture and the mixture was stirred for 30 minutes. Aqueous sodium citrate was then added to the mixture under vacuum and the mixture stirred for a further 30 minutes.

Tests were conducted to measure the ability of high surface area zinc oxide to absorb sebum.

- 5 Unless otherwise stated, the zinc oxide used in the following examples has an average surface area of 90m²/g and an average particle diameter of 10.47 μ m.

This method was taken from American Society For Testing and Materials, 1962, Standard Method of Test for Oil Absorption of Pigments by Spatula Rub-Out.
10 ASTM Standards 1961, p213-214.

The recipe for the artificial sebum used was from a paper called "An Original Procedure for Quantitation of Cutaneous Reabsorption of Sebum" by D Blane, D Saint-Leger, J Bandt, S Constans and P Agache, Arch Dermatol Res, 1989,
15 281: p346-350:-

	Palmitic Acid	5.000g
	Squalene	15.000g
	Cholesterol	6.000g
20	Oleic Acid	25.000g
	Lauric Acid	1.000g
	Dervacid 3155	3.000g
	Myristic Acid	3.000g
	Evening Primrose Oil deodorised	34.000g
25	Glycerol Isostearate	3.000g
	Coronet Lanolin	5.000g
<hr/>		
	Total	100.000g

- 30 These were added together and then heated to form the artificial sebum.
1g of high surface area zinc oxide powder was weighed onto a piece of glass.
A beaker of artificial sebum and a pipette were weighed. Artificial sebum was added to the high surface area zinc oxide drop by drop, and mixed thoroughly

after each drop was added. The test was complete when exactly enough sebum was added to the high surface area zinc oxide to produce a very stiff, putty like paste, which did not break or separate. The beaker and pipette were weighted to determine the amount of sebum used. This was then repeated.

5 This was done again using zinc oxide BP and Sanitised talc as controls.

Table 1

10 Table to Show the Sebum Absorbency of High Surface Area Zinc Oxide Compared to Zinc Oxide BP

High Surface Area Zinc Oxide (g)	ZnO BP (g)
Amount of Sebum Absorbed	Amount of Sebum Absorbed
1.41	0.57
1.10	0.60
1.25	0.45
1.24	0.53
1.25	0.67
1.22	0.64
1.67	0.37
1.22	0.57
1.45	0.56
1.32	0.56
Averages	Averages
1.31	0.55

Expressed as a % difference *, these results show that high surface area zinc oxide absorbs an average 76% more artificial sebum than zinc oxide BP

- 15 * % difference is the difference between the amount of sebum absorbed by high surface area zinc oxide and the amount absorbed by zinc oxide BP, relative to the amount of sebum absorbed by zinc oxide BP.

Table 2

Table to Show the Sebum Absorbency of High Surface Area Zinc Oxide Compared to Sanitised Talc

5

High Surface Area Zinc Oxide (g)	Sanitised Talc
Amount of Sebum Absorbed	Amount of Sebum Absorbed
1.41	0.84
1.10	0.81
1.25	0.72
1.24	0.76
1.25	0.92
1.22	0.87
1.67	0.64
1.22	1.59
1.45	0.85
1.32	0.94
Averages	Averages
1.31	0.89

Expressed as a % difference *, these results show that high surface area zinc oxide absorbs on average 47% more artificial sebum than sanitised talc.

10

* % difference is the difference between the amount of sebum absorbed by high surface area zinc oxide and the amount absorbed by talc, relative to the amount of sebum absorbed by talc.

Overall the high surface area zinc oxide is better at absorbing artificial sebum than Zinc Oxide BP and Sanitised Talc.

15

Sebum Fatty Acid Absorption

Sebum fatty acid absorption by high surface area zinc oxide was analysed using gas chromatography.

Method

- 5 The artificial sebum was prepared for analysis in the following way: 1% of high surface area zinc oxide was put into a beaker with 99% of artificial sebum. This was mixed for half an hour with a magnetic stirrer, to make sure that the high surface area zinc oxide was fully mixed in and in contact with all of the sebum. The solution was then centrifuged for two minutes at 14,000 rpm. The
- 10 supernatant was removed and samples of this were sent for fatty acids analysis using gas chromatography. This was repeated for a mixture containing 1.33% high surface area zinc oxide and 98.66% artificial sebum. A sample of artificial sebum which had not had any high surface area zinc oxide added to it was used as a control.

Table 3

Table to Show the Absorption of Specific Fatty Acids by the High Surface Area Zinc Oxide

5

Fatty Acid	Chain Length	Sebum only (control)	Sebum With 1% ZnO S	Percentage Fatty Acid Absorbed	Sebum with 1.33% ZnO S	Percentage Fatty Acid Absorbed
% Lauric Acid	C12	0.95	0.90	5.26	0.93	2.11
% Myristic Acid	C14	3.17	3.11	1.89	2.81	11.36
% Palmitic Acid	C16	6.17	5.20	15.72	5.10	17.34
% Stearic Acid	C18	3.67	3.07	16.35	3.08	16.08
% Oleic Acid	C18, with 1 unsaturated bond	28.60	25.10	12.24	25.30	11.54

These results show that high surface area zinc oxide preferentially absorbs these longer chain fatty acids. Longer chain fatty acids are more comedogenic than shorter chain fatty acids.

10

Tests were conducted to measure the ability of high surface area zinc oxide to absorb sweat, in a similar to that described above for sebum absorption.

Method

This method was taken from American Society for Testing and Materials, 1962. Standard Method of Test for Oil Absorption of Pigments by Spatula Rub-

Out. ASTM Standards 1961, p213-214 and modified for measuring sweat absorption.

5 The recipe for the artificial sweat was taken and modified from Geigy Scientific Tablets. Units of Measurements. Body Fluids. Composition of The Body. Nutrition. C Lentner. Ciba-Geigy. 1981. p108-112:

	Water	994.060g
	Urea	1.990g
	Glucose	0.170g
10	Lactic Acid	1.190g
	Leucine	2.590g
<hr/>		
	Total	1000.000g

15 The above raw materials were dissolved together to form the artificial sweat.

Table 4

5 Table to Show the Sweat Absorbency of High Surface Area Zinc Oxide Compared to Zinc Oxide BP

High Surface Area ZnO (g)	ZnO BP (g)
Amount of Sweat Absorbed	Amount of Sweat Absorbed
2.90	0.60
1.40	0.50
2.00	0.60
1.60	0.60
1.40	0.20
1.60	0.50
1.30	1.10
1.50	0.70
1.20	0.50
1.50	0.50
Averages	Averages
1.64	0.58

10 Expressed as a % difference *, these results show that high surface area zinc oxide absorbs on average 183% more artificial sweat than zinc oxide BP

Thus these results show that high surface area zinc oxide absorbs artificial sweat better than Zinc Oxide BP.

* % difference is the difference between the amount of sweat absorbed by high surface area zinc oxide and the amount absorbed by zinc oxide BP, relative to the amount of sweat absorbed by zinc oxide BP.

5 **Table 5**

Table to Show the Sweat Absorbency of High Surface Area Zinc Oxide Compared to Sanitised Talc

10

High Surface Area ZnO (g)	Sanitised Talc (g)
Amount of Sweat Absorbed	Amount of Sweat Absorbed
2.90	1.00
1.40	1.70
2.00	1.10
1.60	0.70
1.40	0.90
1.60	0.90
1.30	0.90
1.50	0.90
1.20	1.10
1.50	0.80
Averages 1.64	Averages 1.0

Expressed as a % difference *, these results show that on average, high surface area zinc oxide absorbs 64% more artificial sweat than sanitised talc.

Thus these results show that high surface area zinc oxide absorbs artificial sweat better than sanitised talc.

- * % difference is the difference between the amount of sweat absorbed by high surface area zinc oxide and the amount absorbed by talc, relative to the amount of sweat absorbed by the talc.

Tests were conducted to measure the ability of foot powder formulations to absorb water

- 1g of foot powder containing high surface area zinc oxide powder (as made in Example 3) was weighed onto a piece of glass. Water was added to the foot powder drop by drop, and mixed thoroughly after each drop was added. The test was completed when exactly enough water was added to the foot powder to produce a very stiff, putty like paste, which did not break or separate. The beaker and pipette were weighed to determine the amount of water used. This was then repeated using a control of a standard foot powder (as made in Example 2).

TABLE 6

Table to show the absorbency of foot powder with high surface area zinc oxide compared to standard without high surface area zinc oxide

5

Water absorbed by foot powder containing high surface area zinc oxide (g)	Water absorbed by standard foot powder (g)
1.17	1.13
1.26	0.94
1	0.97
1.11	0.96
1.2	0.97
1.15	1.13
1.1	1.02
1.21	0.97
1.05	0.98
1.12	0.87
Average 1.37	Average 0.994

Expressed as a % difference *, these results show that on average, foot powder containing high surface area zinc oxide absorbs 15% more water than the standard foot powder.

10

Thus these results show that foot powder containing high surface area zinc oxide is more efficacious at absorbing water than the conventional foot powder formulation.

15

* % difference is the difference between the amount of water absorbed by foot powder containing high surface area zinc oxide, and the amount absorbed by standard foot powder, relative to the amount absorbed by standard foot powder.

CLAIMS

1. Topical composition comprising a cosmetically acceptable diluent or carrier and high surface area zinc oxide, having a surface area between 30m²/g and 100m²/g and a particle size between 0.1 and 200 μ m in diameter, said high surface area zinc oxide being present in an amount sufficient to absorb liquids from the parts of the body to which the topical composition is applied.
5
2. A composition as claimed in claim 1 where the surface area of the high surface area zinc oxide is between 90 and 100m²/g and the particle size is between 0.1 and 20.5 μ m in diameter.
10
3. A composition as claimed in claim 1 where the high surface area zinc oxide is present from 1 to 10% by weight of the total composition.
4. A composition as claimed in claim 1 where the high surface area zinc oxide is present from 3 to 8% by weight of the total composition.
5. A composition as claimed in claim 1 where the high surface area zinc oxide is present from 4 to 6% by weight of the total composition.
15
6. The use of high surface area zinc oxide to absorb liquids from the body where the liquids are sweat and/or sebum.
7. The use of high surface area zinc oxide in the treatment of acne, athletes foot and nappy rash.
20
8. The use of high surface area zinc oxide in the preparation of a medicament for the treatment of acne, athletes foot and nappy rash.

INTERNATIONAL SEARCH REPORT

International Application No

PCT/EP 99/03590

A. CLASSIFICATION OF SUBJECT MATTER

IPC 6 A61K7/48 A61K7/36 A61K33/30

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 6 A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	PATENT ABSTRACTS OF JAPAN vol. 1996, no. - , 26 December 1996 (1996-12-26) & JP 08 217637 A (KAO CORP), 27 August 1996 (1996-08-27) abstract	1-6,8
P,X	PATENT ABSTRACTS OF JAPAN vol. 1999, no. 053, 31 May 1999 (1999-05-31) & JP 11 049637 A (KAO CORP), 23 February 1999 (1999-02-23) abstract	1-6,8
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☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

* Special categories of cited documents :

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- "P" document published prior to the international filing date but later than the priority date claimed

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- "&" document member of the same patent family

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INTERNATIONAL SEARCH REPORT

International Application No

PCT/EP 99/03590

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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